



FORM PT-1449

U.S. DEPARTMENT OF COMMERCE  
PATENT AND TRADEMARK OFFICEINFORMATION DISCLOSURE  
STATEMENT BY APPLICANTATTY. DOCKET NO.  
Cell 4.8 FWC CPASERIAL NO.  
08/923,138APPLICANT  
Raju Kucherlapati et al.CONFIRMATION NO.  
2306FILING DATE  
September 4, 1997GROUP  
1632

## OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

EXAMINER INITIAL	
NO	Albertsen, et al., Construction and characterization of a yeast artificial chromosome library containing seven haploid human genome equivalents," <i>Proc. Natl. Acad. Sci.</i> , 87:4256-4260 (1990).
	Aldhous, "Transgenic mice display a class (switching) act," <i>Science</i> 262:1212-1213 (1993).
	Ayares, et al., "Sequence homology requirements for intermolecular recombination in mammalian cells," <i>Proc. Natl. Acad. Sci.</i> , 83:5199-5203 (1986).
	Berman, et al., "Content and organization of the human Ig V <sub>H</sub> locus: definition of three new V <sub>H</sub> families and linkage to the Ig C <sub>H</sub> locus," <i>EMBO J.</i> , 7:727-738 (1988).
	Bird, et al., "Single-Chain Antigen-Binding Proteins," <i>Science</i> , 243:423-426 (1988).
	Blankenstein, et al., "Immunoglobulin V <sub>H</sub> region genes of the mouse are organized in overlapping clusters," <i>Eur. J. Immunol.</i> , 17:1351-1357 (1987).
	Brinster, et al., "Introns increase transcriptional efficiency in transgenic mice," <i>Proc. Natl. Acad. Sci.</i> , 85:836-840 (1988).
	Brownstein, et al., "Isolation of single-copy human genes from a library of yeast artificial chromosome clones," <i>Science</i> , 244:1348-1351 (1989).
	Brüggemann, et al., "Human antibody production in transgenic mice: expression from 100kb of the human IgH locus," <i>European Journal of Immunology</i> , 21:1323-1326 (1991).
	Brüggemann, et al., "Construction, function and immunogenicity of recombinant monoclonal antibodies," <i>Behring Inst. Mitt.</i> , 87:21-24 (1990).
	Burke, et al., "Cloning of large segments of exogenous DNA into yeast by means of artificial chromosome vectors," <i>Science</i> , 236:806-812 (1987).
	Buttin, G., "Exogenous Ig gene rearrangement in transgenic mice: a new strategy for human monoclonal antibody production?," <i>Trends in Genetics</i> , 3(8):205-206 (1987).
	Capecchi, et al., "Altering the Genome by Homologous Recombination," <i>Science</i> , 244(16):1288-1292 (1989).
	Choi, et al., "Transgenic mice containing a human heavy chain immunoglobulin gene fragment cloned in a yeast artificial chromosome," <i>Nature Genetics</i> 4:117-123 (1993).
	Choi et al., "RNA Splicing Generates a Variant Light Chain from an Aberrantly Arranged $\kappa$ Gene," <i>Nature</i> , 286: 776-779 (1980).
Davies, et al., "Targeted alterations in yeast artificial chromosomes for inter-species gene transfer," <i>Nuc. Acids Res.</i> , 20:2693-2698 (1992).	
Doelker, et al., "The CySF-L2 factor from dialysable human leucocyte extract activates natural killer cytotoxicity by induction of interferon $\gamma$ ," <i>Cancer Immunology Immunotherapy</i> , 34:299-305 (1992).	

EXAMINER

DATE CONSIDERED

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not conformance and not considered. Include copy of this form with next communication to applicant.



FORM PTO-1499 U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE  INFORMATION DISCLOSURE STATEMENT BY APPLICANT	ATTY. DOCKET NO. Cell 4.8 FWC CPA	SERIAL NO. 08/923,138
	APPLICANT Raju Kucherlapati et al.	CONFIRMATION NO. 2306
	FILING DATE September 4, 1997	GROUP 1632

## OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

EXAMINER INITIAL	
WJ	Doetschman et al., "Targeted Mutation of the Hprt Gene in Mouse Embryonic Stem Cells," <i>Proc. Natl. Acad. Sci. USA</i> , 85: 8583-8587 (1988).
	Eisen, Herman N., "Immunology: An Introduction to Molecular and Cellular Principles of the Immune Responses," 349-351 (2d ed. 1989).
	Eliceiri, et al., "Stable integration and expression in mouse cells of yeast artificial chromosomes harboring human genes," <i>Proc. Natl. Acad. Sci.</i> , 88:2179-2183 (1991).
	Emery, et al., "Humanised monoclonal Antibodies for Therapeutic Applications," <i>Expert Opinion on Investigational Drugs</i> , 3:241-251 (1994).
	Garza, et al., "Mapping the Drosophila genome with yeast artificial chromosomes," <i>Science</i> , 246:641-646 (1989).
	Gnirke, et al., "Cloning and <i>in vivo</i> expression of the human GART gene using yeast artificial chromosomes," <i>EMBO</i> , 10(7):1629-1634 (1991).
	Green et al., "Antigen-specific Human Monoclonal Antibodies from Mice Engineered with Human Ig Heavy and Light Chain Yacs," <i>Nature Genet.</i> , 7:13 (1994).
	Griffiths, et al., "Isolation of high affinity human antibodies directly from large synthetic repertoires," <i>The EMBO Journal</i> , 13:3245-3260 (1994).
	Huxley, et al., "The Human HPRT gene on a yeast artificial chromosome is functional when transferred to mouse cells by cell fusion," <i>Genomics</i> , 9:742-750 (1991).
	Jakobovits et al., "Germ-line Transmission and Expression of a Human-derived Yeast Artificial Chromosome," <i>Nature</i> , 362:252-258 (1993).
	James, et al., "Human monoclonal antibody production: current status and future prospects," <i>Journal of Immunological Methods</i> , 100:5-40 (1987).
	Johnson et al., "Targeting of Nonexpressed Genes in Embryonic Stem Cells via Homologous Recombination," <i>Science</i> , 245:1234-1236 (1989).
	Joyner, et al., "Production of a mutation in mouse En-2 gene by homologous recombination in embryonic stem cells," <i>Nature</i> , 338:153-155 (1989).
	Koller, et al., "Inactivating the $\beta_2$ -microglobulin locus in mouse embryonic stem cells by homologous recombination," <i>Proc. Natl. Acad. Sci.</i> , 86:8932-8935 (1989).
	Kucherlapati, R., "Homologous recombination in mammalian somatic cells," <i>Prog. Nucleic Acid Res. Mol. Biol.</i> , 36:301-310 (1989).

RECEIVED

OCT 02 2002

TECH CENTER 1600/2900

EXAMINER

DATE CONSIDERED

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not conformance and not considered. Include copy of this form with next communication to applicant.

U.S. DEPARTMENT OF COMMERCE  
PATENT AND TRADEMARK OFFICEINFORMATION DISCLOSURE  
STATEMENT BY APPLICANTATTY. DOCKET NO.  
Cell 4.8 FWC CPASERIAL NO.  
08/923,138APPLICANT  
Raju Kucherlapati et al.CONFIRMATION NO.  
2306FILING DATE  
September 4, 1997GROUP  
1632

## OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

EXAMINER INITIAL	
<i>AS</i>	Lenz, et al., "Expression of heterobispecific antibodies by genes transferred into producer hybridoma cells," <i>Gene</i> , 87:213-218 (1990).
	Liu et al., "Chimeric mouse-human IgG1 antibody that can mediate lysis of cancer cells," <i>Proc Natl Acad Sci USA</i> , 84:3439-3443 (1987).
	Mansour, et al., "Disruption of the proto-oncogene int-2 in mouse embryo-derived stem cells: a general strategy for targeting mutations to non-selectable genes," <i>Nature</i> , 336:348-352 (1988).
	Matsuda, et al., "Structure and physical map of 64 variable segments in the 3' 0.8- megabase region of the human immunoglobulin heavy-chain locus," <i>Nature Genetics</i> , 3:88-94 (1993).
	Max et al., "Sequences of Five Potential Recombination Sites Encoded Close to an Immunoglobulin K Constant Region Gene," <i>Proc. Natl. Acad. Sci. USA</i> , 76:3450-3454 (1979).
	Miller, et al., "Structural alterations in J regions of mouse immunoglobulin lambda genes are associated with differential gene expression," <i>Nature</i> , 295:428-430 (1982).
	Morrison et al., "Success Is in the Specification," <i>Nature</i> , 368:812-813 (1994).
	Mortensen, et al., "Production of homozygous mutant ES cells with a single targeting construct," <i>Mol. Cell. Biol.</i> , 12(5):2391-2395 (1992).
	Munker, et al., "Recombinant human TNF induces production of granulocyte-monocyte colony-stimulating factor," <i>Nature</i> , 323:79-82 (1986).
	Orkin et al., "Mutation in an Intervening Sequence Splice Junction in Man," <i>Proc. Natl. Acad. Sci. USA</i> , 78:5041-5045 (1981).
	Pachnis, et al., "Transfer of a yeast artificial chromosome carrying human DNA from <i>Saccharomyces cerevisiae</i> into mammalian cells," <i>Proc. Natl. Acad. Sci.</i> , 87:5109-5113 (1990).
	Pavan, et al., "Modification and transfer into an embryonal carcinoma cell line of a 360-kilobase human-derived yeast artificial chromosome," <i>Mol. Cell. Biol.</i> , 10(8):4163-4169 (1990).
	Queen, et al., "A humanized antibody that binds to the interleukin 2 receptor," <i>Proc. Natl. Acad. Sci. USA</i> , 86:10029-10033 (1989).
	Rajewsky et al., "Evolutionary and Somatic Selection of the Antibody Repertoire in the Mouse," <i>Science</i> , 238:1088-1094 (1987).
	Ramirez-Solis et al., "Chromosome Engineering in Mice," <i>Nature</i> , 378:720-724 (1995).
	Sakano, et al., "Identification and nucleotide sequence of a diversity DNA segment (D) of immunoglobulin heavy-chain genes," <i>Nature</i> , 290:562-565 (1981).

RECEIVED

OCT 02 2002

TECH CENTER 1600/2900

EXAMINER

*AS*

DATE CONSIDERED

*10/15*

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not conformance and not considered. Include copy of this form with next communication to applicant.

7

U.S. DEPARTMENT OF COMMERCE  
PATENT AND TRADEMARK OFFICEINFORMATION DISCLOSURE  
STATEMENT BY APPLICANTATTY. DOCKET NO.  
Cell 4.8 FWC CPASERIAL NO.  
08/923,138APPLICANT  
Raju Kucherlapati et al.CONFIRMATION NO.  
2306FILING DATE  
September 4, 1997GROUP  
1632

## OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

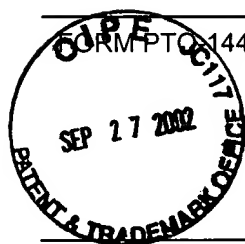
EXAMINER INITIAL	
<i>[Signature]</i>	Sakano, et al., "Two Types of Somatic Recombination are Necessary for the Generation of Complete Immunoglobulin Heavy-Chain Genes," <i>Nature</i> 286:676-683 (1980).
	Sakano et al., "Sequences at the Somatic Recombination Sites of Immunoglobulin Light-chain Genes," <i>Nature</i> , 280:288-294 (1979).
	Schedl, et al., "A Method for the Generation of YAC Transgenic Mice by Pronuclear Microinjection," <i>Nucleic Acids Research</i> 21(20):4783-4787 (1993).
	Schedl, et al., "A yeast artificial chromosome covering the tyrosinase gene confers copy number-dependent expression in transgenic mice," <i>Nature</i> , 362:258-261 (1993).
	Schedl et al., "Transgenic Mice Generated by Pronuclear Injection of a Yeast Artificial Chromosome," <i>Nucleic Acids Res.</i> , 20:3073-3077 (1992).
	Schwartzberg et al., "Germ-line Transmission of a c-abl Mutation Produced by Targeted Gene Disruption in ES Cells," <i>Science</i> , 246:799-803 (1989).
	Seidman, et al., "A Mutant Immunoglobulin Light Chain Is Formed by Aberrant DNA- and RNA- Splicing Events," <i>Nature</i> , 286:779-783 (1980).
	Shimizu, et al., "Immunoglobulin double-isotype expression by trans-mRNA in a human immunoglobulin transgenic mouse," <i>Proc. Natl. Acad. Sci.</i> , 86:8020-8023 (1989).
	Shin, et al., "Physical map of the 3' region of the human immunoglobulin heavy chain locus: clustering of autoantibody-related variable segments in one haplotype," <i>EMBO J</i> , 10:3641-3645 (1991).
	Strauss, et al., "Germ Line Transmission of a Yeast Artificial Chromosome Spanning the Murine $\alpha_1(1)$ Collagen Locus," <i>Science</i> 259:1904-1907 (1993).
	Thomas, et al., "Site-directed metagenesis by gene targeting in mouse embryo-derived stem cells," <i>Cell</i> , 51:503-512 (1987).
	Traver, et al., "Rapid screening of a human genomic library in yeast artificial chromosomes for single-copy sequences," <i>Proc. Natl. Acad. Sci.</i> , 86:5898-5902 (1989).
	Treisman et al., "Specific Transcription and RNA Splicing Defects in Five Cloned $\beta$ -thalassaemia Genes," <i>Nature</i> , 302:591-596 (1983).
	Tucker, et al., "Mouse IgA heavy chain gene sequence: implications for evolution of immunoglobulin hinge exons," <i>Proc. Natl. Acad. Sci.</i> , 78:7684-7688 (1981).
	Yancopoulos, et al. "Developmentally Controlled and Tissue-Specific Expression of Unrearranged $V_H$ Gene Segments," <i>Cell</i> 40:271-281 (1985)
	Zachau, "The human immunoglobulin $\kappa$ locus and some of its acrobatics," <i>Biol. Chem.</i> , 371:1-6 (1990)

EXAMINER

DATE CONSIDERED

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not conformance and not considered. Include copy of this form with next communication to applicant.

8



U.S. DEPARTMENT OF COMMERCE  
PATENT AND TRADEMARK OFFICE

INFORMATION DISCLOSURE  
STATEMENT BY APPLICANT

ATTY. DOCKET NO.  
Cell 4.8 FWC CPA

SERIAL NO.  
08/923,138

APPLICANT  
Raju Kucherlapati et al.

CONFIRMATION NO.  
2306

FILING DATE  
September 4, 1997

GROUP  
1632

OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

EXAMINER INITIAL	
<i>10</i>	Zijlstra et al., "Germ-line Transmission of a Disrupted $\beta$ 2-microglobulin Gene Produced by Homologous Recombination in Embryonic Stem Cells," <i>Nature</i> , 342:435-438 (1989)

RECEIVED

OCT 02 2002

TECH CENTER 1600/2900

EXAMINER

*Dule*

DATE CONSIDERED

*10/16/02*

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not conformance and not considered. Include copy of this form with next communication to applicant.